Application No. 10/536,569

Amendment dated April 13, 2010

Docket No.: 1258\_3378US

Reply to Office Action of October 14, 2009

## AMENDMENTS TO THE CLAIMS

This listing of claims will replace all prior versions, and listings, of claims in the application:

Claims:

Claims 1 - 39. (Canceled)

Claim 40. (Currently amended) A fluorescent silica-based nanoparticle comprising:

a silica-based core comprising <u>and organic functional group</u> comprising a mercapto <u>substituent</u> <del>group</del>;

[[a]] an organic fluorescent compound positioned within the silicabased core; and

a silica shell surrounding at least a portion of the core,

wherein the fluorescent silica-based nanoparticle eemprises has a diameter between about 10 nm and about 70 nm of 70 nm or less and is conjugated to a ligand.

Claims 41-44. (Canceled)

Claim 45. (New) The fluorescent silica-based nanoparticle of claim 40, wherein the diameter is 50 nm or less.

Claim 46. (New) The fluorescent silica-based nanoparticle of claim 40, wherein the ligand comprises at least one of a protein, a peptide, or an oligopeptide.

Claim 47. (New) The fluorescent silica-based nanoparticle of claim 40, wherein the ligand is conjugated to the nanoparticle by a coupling agent comprising at least one carbon containing linkage, the carbon containing linkage selected from the group comprising an ester linkage, a thiolester linkage, an amide linkage, a sulfate ester linkage and combinations thereof.

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Claim 48. (New) The fluorescent silica-based nanoparticle of claim 40, wherein the silica shell covers from about 10 percent to about 100 percent of the surface area of the core.

- Claim 49. (New) The fluorescent silica-based nanoparticle of claim 40, wherein the silica-based core further comprises a silica-based network, wherein the fluorescent compound is covalently attached to the silica-based network.
- Claim 50. (New) The fluorescent silica-based nanoparticle of claim 40, wherein the fluorescent quantum yield of the fluorescent organic dye in the nanoparticle is about two-fold to about three-fold greater than the fluorescent quantum yield of the same fluorescent organic dye free in aqueous solution.
- Claim 51. (New) The fluorescent silica-based nanoparticle of claim 40, the mercapto substituent is bonded to a maleimide.
- Claim 52. (New) The fluorescent silica-based nanoparticle of claim 40, wherein the silica-based core has a radius between about 1.6 nm and about 3.5 nm.
- Claim 53. (New) The fluorescent silica-based nanoparticle of claim 40, wherein the silica-based core has a radius between about 2.2 nm and about 2.9 nm.